WHAT IS CLAIMED IS:

1. An amyloid binding compound having one of structures A-E or a water soluble, non-toxic salt thereof:



wherein Z is S, NR', O or CR' in which case the/correct tautomeric form of the heterocyclic ring becomes an indole in which R' is H or a lower alkyl group:

wherein Y is NR¹R², OR², or SR²;

$$Z$$
or
 R'

wherein the nitrogen of amine;

is not a quaternary

or an amyloid binding compound having one of structures F-J or a water soluble, non-toxic salt thereof:

Structure F
$$R_{13}$$
 R_{14} R_{14} R_{15} R_{10} R_{10}

wherein each Q is independently selected from one of the following structures:



$$R_6$$
 R_5 wherein $n = 0, 1, 2, 3 \text{ or } 4,$
 R_4 R_3 R_4 R_5 R_5 R_6 R_5 R_6 R_7 R_8 R_8 R_8 R_8 R_8 R_8 R_8 R_8 R_8 R_8

wherein Z is S, NR', O, or C(RY)2 in which R' is H or a lower alkyl group; wherein U is CR' (in which R' is H or a lower alkyl group) or N (except when U

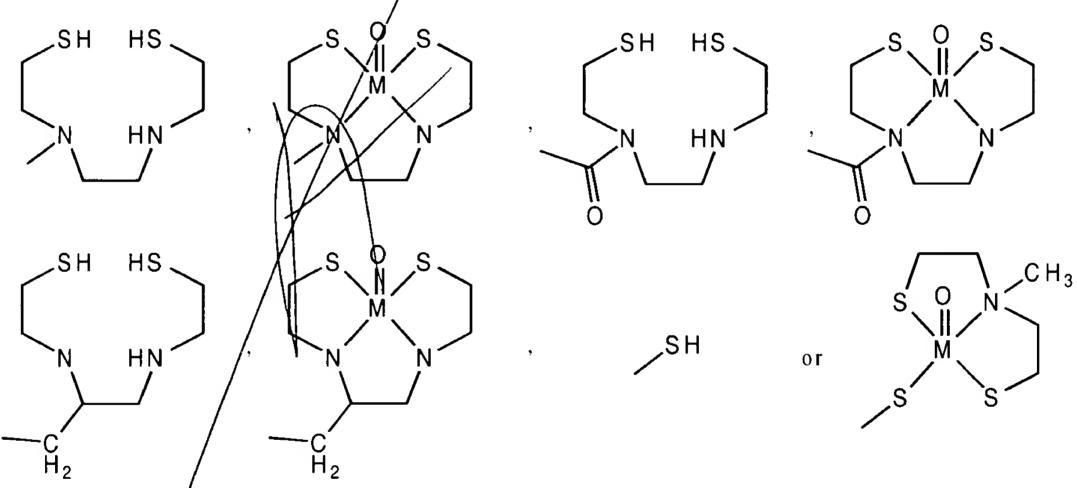
$$R_6$$
 R_5 Y R_4 R_3 Y

= N, then Q is not $\frac{R_4}{N} \setminus \frac{R_3}{N}$ wherein Y is NR¹R², O,R², or SR²;

amine;

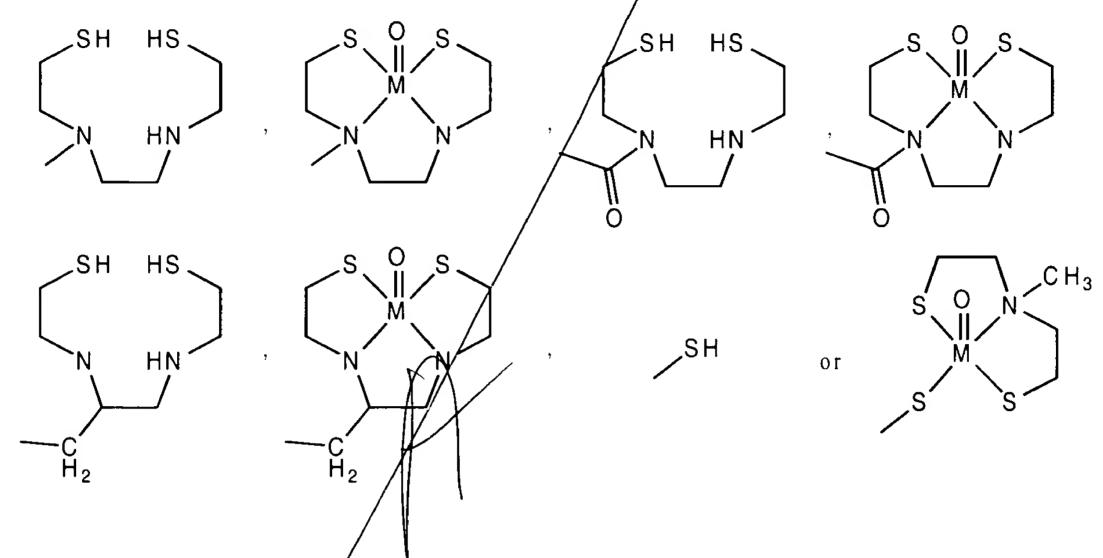
wherein each R^1 and R^2 independently is selected from the group consisting of H, a lower alkyl group, $(CH_2)_nOR'$ (wherein n=1, 2, or 3), CF_3 , CH_2-CH_2X , CH_2-CH_2X (wherein X=F, CI, Br or I), (C=O)-R', R_{ph} , and $(CH_2)_nR_{ph}$ (wherein n=1, 2, 3, or 4 and R_{ph} represents an unsubstituted or substituted phenyl group with the phenyl substituents being chosen from any of the non-phenyl substituents defined below for R^3-R^{14} and R' is H or a lower alkyl group);

and wherein each R^3 - R^{14} independently is selected from the group consisting of H, F, CI, Br, I, a lower alkyl group, (CH₂)_nOR' (wherein n = 1, 2, or 3), CF₃, CH₂-CH₂X, O-CH₂-CH₂X, CH₂-CH₂X, O-CH₂-CH₂CH₂CH₂CH₂X (wherein X = F, CI, Br or I), CN, (C = O)-R', N(R')₂, NO₂, (C = O)N(R')₂, O(CO)R', OR', SR', COOR', R_{ph}, CR' = CR'-R_{ph}, CR₂'-CR₂'-R_{ph} (wherein R_{ph} represents an unsubstituted or substituted phenyl group with the phenyl substituents being chosen from any of the non-phenyl substituents defined for R¹-R¹⁴ and wherein R' is H or a lower alkyl group), a tri-alkyl tin and a chelating group (with or without a chelated metal group) of the form W-L or V-W-L, wherein V is selected from the group consisting of -COO-, -CO-, -CH₂O- and -CH₂NH-; W is -(CH₂)_n where n = 0,1,2,3,4, or 5; and L is:



wherein M is/selected from the group consisting of Tc and Re;

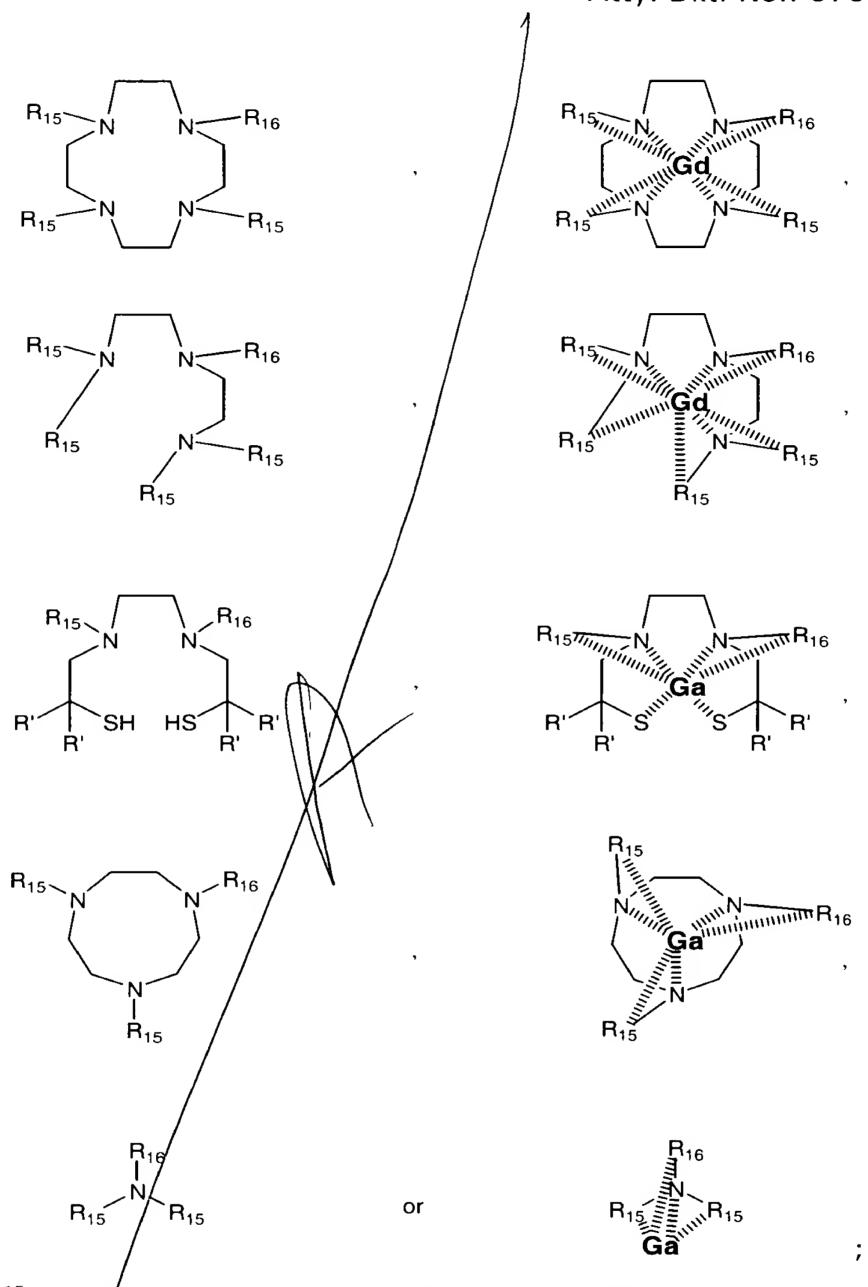
or wherein each R^1 and R^2 is a chelating group (with or without a chelated metal group) of the form W-L, wherein W is $-(CH_2)$, where n=2,3,4, or 5; and L is:



wherein M is selected from the group consisting of Tc and Re; or wherein each R^1 – $R^{1/4}$ independently is selected from the group consisting of a chelating group (with or without a chelated metal ion) of the form W-L and V-W-L, wherein V is selected from the group consisting of –COO-, and -CO-; W is – (CH₂)_n where n = 0,1,2,3,4, or 5; L is:

and wherein R¹⁵ independently is selected from the following:

or an amyloid binding, chelating compound (with or without a chelated metal group) or a water soluble, non-toxic salt thereof of the form:



wherein R¹⁵ independently is selected from the following:

H,
$$COOH$$
. $CONHCH_3$. CH_3



and
$$R^{16}$$
 is R_{23} R_{24} R_{17} R_{18} R_{18} R_{20} R_{20} R_{20} R_{20} R_{20} , wherein Q is

independently selected from one of the following structures:

R₁₇ R₁₈ wherein
$$n = 0, 1, 2, 3 \text{ or } 4,$$

R₁₇ R₁₈

R₁₇ R₁₈

R₁₇ R₁₈

R₁₈ R₁₉

R₁₇ R₁₈

R₁₇ R₁₈

R₁₈ R₁₉

R₁₉ R₁₉

R₁₀ R₁₉

wherein Z is S, NR', O, or $C(R)_2$ in which R' is H or a lower alkyl group; wherein U is N or CR'; wherein Y is NR^1R^2 , OR^2 , or SR^2 ;

wherein each R^{17} - R^{24} independently is selected from the group consisting of H, F, CI, Br, I, a lower alkyl group, $(CH_2)_nOR'$ (wherein n=1, 2, or 3), CF_3 , CH_2 - CH_2X , $O-CH_2$ - CH_2X , CH_2 - CH_2X , $O-CH_2$ - CH_2 - CH_2X (wherein X=F, CI, Br or I), CN, (C=O)-R', $N(R')_2$, NO_2 , $(C=O)N(R')_2$, O(CO)R', OR', SR', COOR', R_{ph} , $CR'=CR'-R_{ph}$ and $CR_2'-CR_2'-R_{ph}$ (wherein R_{ph} represents an unsubstituted or substituted phenyl group with the phenyl substituents being chosen from any of the non-phenyl substituents defined for $R^{17}-R^{20}$ and wherein R' is H or a lower alkyl group).

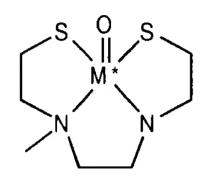
2. The compound of claim 1, wherein at least one of the substituents R¹-R¹⁴ is selected from the group consisting of ¹³¹I, ¹²³I, ¹⁶Br, ¹⁶Br, ¹⁶Br, ¹⁶Br, ¹⁶Br, ¹⁶Br, ¹⅙F, CH₂-CH₂-X*, O-

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Atty. Dkt. No.: 076333-0281

CH₂-CH₂-X*, CH₂-CH₂-CH₂-X*, O- CH₂-CH₂-CH₂-X* (wherein X* = 131 I, 123 I, 76 Br, 75 Br or 18 F), 19 F, 125 I, a carbon-containing substituent as specified in claim 1 wherein at least one carbon is 11 C or 13 C and a chelating group (with chelated metal group) of the form W-L* or V-W-L*, wherein V is selected from the group consisting of -COO-, -CO-, -CH₂O- and -CH₂NH-; W is -(CH₂)_n where

n = 0, 1, 2, 3, 4, or 5; and L* is:

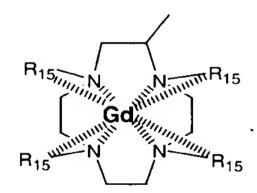


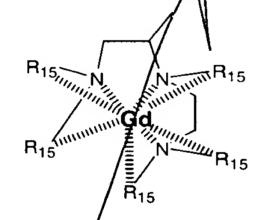
or

or

wherein M* is 99mTc;

and a chelating group (with chelated metal group) of the form W-L* or V-W-L*, wherein V is selected from the group consisting of $-COO_{-}$, $-CO_{-}$, $-CH_{2}O_{-}$ and $-CH_{2}NH_{-}$; W is $-(CH_{2})_{n}$ where n=0,1,2,3,4, or 5; and L* is:





and wherein R¹⁵ independently is selected from the following:

H, COOH, CONHCH₃. CH₃. SH . HS or
$$\stackrel{\text{HO}}{\longrightarrow}$$
 ;

or the chelating compound of claim 1 (with chelated metal group) of the form:

wherein R¹⁵ independently is selected from the following:

H, COOH. CONHCH
$$_3$$
. OH SH $_2$ R $_{17}$ R $_{18}$ CH $_2$ and R $_{16}$ is $_{16}$ is $_{18}$ Or $_{$

independently selected from one of the following structures:

wherein
$$n = 0, 1, 2, 3 \text{ or } 4,$$

$$R_{10}$$

$$R_{11}$$

$$R_{11}$$

$$R_{12}$$

$$R_{11}$$

$$R_{12}$$

$$R_{13}$$

$$R_{14}$$

$$R_{15}$$

$$R_{15}$$

$$R_{19}$$

$$R_{19}$$

$$R_{19}$$

wherein Z is S, NR', O, or C(R')₂ in which R' is H or a lower alkyl group; wherein U is N or CR';

(wherein X = F, CI, B/OrI);

wherein Y is NR1R2, OR2, or SR2;

wherein each R^{17} - R^{24} independently is selected from the group consisting of H, F, CI, Br, I, a lower alkyl group, $(CH_2)_nOR'$ (wherein n=1, 2, or 3), CF_3 , CH_2 - CH_2X , O- CH_2 - CH_2X , O- CH_2 - CH_2X , O- CH_2 - CH_2X (wherein X=F, CI, Br or I), CN, (C=0)-R', $N(R')_2$, NO_2 , $(C=0)N(R')_2$, O(CO)R', OR', SR', COOR', R_{ph} , CR' = CR'- R_{ph} and CR_2' - CR_2' - R_{ph} (wherein R_{ph} represents an unsubstituted or substituted phenyl group with the phenyl substituents being chosen from any of the non-phenyl substituents defined for R^{17} - R^{20} and wherein R' is H or a lower alkyl group).

3. The compound of claim 1, wherein, Z=S, Y=N, R¹=H; and wherein when the amyloid binding compound of claim 1 is structure A or E, then R² is selected from the group consisting of a lower alkyl group, (CH₂)nOR′ (wherein n = 1, 2, or 3), CF₃, CH₂-CH₂X, CH₂-CH₂-CH₂X (wherein X=F, Cl, Br or l), (C=O)-R′, Rph, and (CH₂)nRph wherein n = 1, 2, 3, or 4; wherein when the amyloid binding compound of claim 1 is structure B, then R² is selected from the group consisting of (CH₂)nOR′ (wherein n=1, 2, or 3, and where when R′=H or CH₃, n is not 1). CF₃, CH₂-CH₂X and CH₂-CH₂-CH₂X

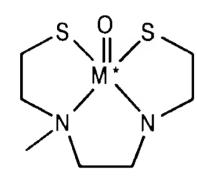
wherein when the amyloid binding compound of claim 1 is structure C, then R^2 is selected from the group consisting of a lower alkyl group, $(CH_2)_nOR'$ (wherein n=1, 2, or 3, CF_3), CH_2-CH_2X , $CH_2-CH_2-CH_2X$ (wherein X=F, CI, Br or I), (C=O)-H, R_p , and $(CH_2)_nR_{ph}$ wherein n=1, 2, 3, or 4; and

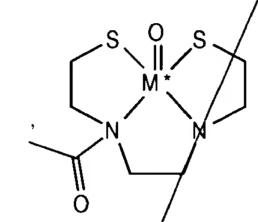
wherein when the amyloid binding compound of claim 1 is structure D, then R^2 is selected from the group consisting of $(CH_2)_nOR'$ (wherein n=1, 2, or 3), CF_3 , CH_2 - CH_2X , CH_2 - CH_2 - CH_2 X (wherein X=F, CI, Br or I), (C=O)-R', R_{ph} , and $(CH_2)_nR_{ph}$ (wherein n=1, 2, 3, or 4) wherein when R^2 is CH_2R_{ph} R^8 is not CH_3 .

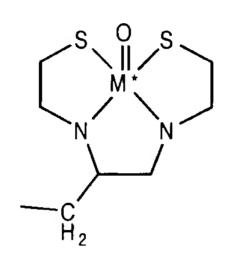
4. The compound of claim 3, wherein at least one of the substituents R³- R¹⁴ is selected from the group consisting of ¹³¹I, ¹²³I, ⁷⁶Br, ⁷⁵Br, ¹⁸F, CH₂-CH₂-X*, O-

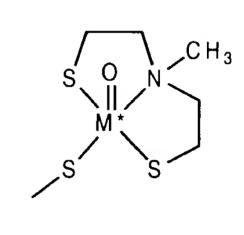


CH₂-CH₂-X*, CH₂-CH₂-CH₂-X*, O- CH₂-CH₂-CH₂-X* (wherein X* = 131 I, 123 I, 76 Br, 75 Br or 18 F), 19 F, 125 I, a carbon-containing substituent as specified in claim 1 wherein at least one carbon is 11 C or 13 C, a chelating group (with chelated metal group) of the form W-L* or V-W-L*, wherein V is selected from the group consisting of $^{-}$ COO-, $^{-}$ CO-, $^{-}$ CH₂O- and $^{-}$ CH₂NH-; W is $^{-}$ (CH₂)_n where $^{-}$ D=0,1,2,3,4, or 5; and L* is:





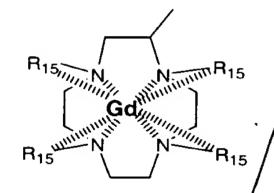


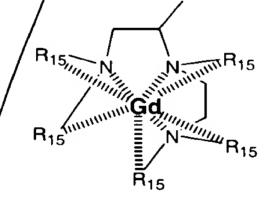


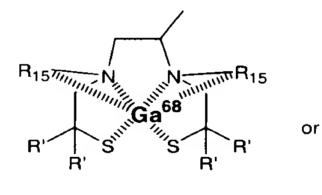
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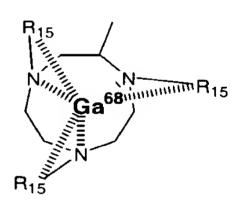
wherein M is 99m Tex

and a chelating group (with chelated metal group) of the form W-L* or V-W-L*, wherein V is selected from the group consisting of $-COO_{-}$, $-CO_{-}$, $-CH_{2}O_{-}$ and $-CH_{2}NH_{-}$; W is $-(CH_{2})_{n}$ where n=0,1,2,3,4, or 5; and L* is:



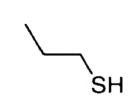


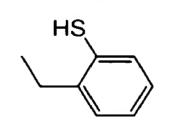




and wherein R¹/₇ independently is selected from the following:

H, COOH, CONHCH₃, CH₃





or the chelating compound of claim 1 (with chelated metal group) of the form:

Harry and harry and harry and harry

wherein R¹⁵ independently/is selected from one of the following structures:

CH₃ COOH ·CONHCH3 , НÓ

, wherein Q is

and R^{16} is

independently selected from one of the following structures:

$$R_{17}$$
 R_{18} $(CH_2)_n$ Y R_{20} R_{19}

wherein n = 0, 1, 2, 3 or 4,

$$R_{17}$$
 R_{18}
 R_{19}
 R_{20}
 R_{19}

$$\begin{array}{c|c}
\hline
 & Z & R_{19} \\
\hline
 & R_{20} & Z & R_{19}
\end{array}$$

wherein Z is S, NR', O, or C(R')2 in which R' is H or a lower alkyl group;

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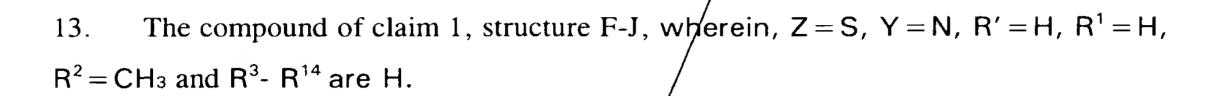
wherein U is N or CR';

wherein Y is NR1R2, OR2, or SR2;

wherein each R^{17} - R^{24} independently is selected from the group consisting of H, F, CI, Br, I, a lower alkyl group, $(CH_2)_nOR'$ (wherein n=1, 2, or 3), CF_3 , CH_2 - CH_2X , O- CH_2 - CH_2X , O- CH_2 - CH_2X , O- CH_2 - CH_2X (wherein X=F, CI, Br or I), CN, (C=O)-R', $N(R')_2$, NO_2 , $(C=O)N(R')_2$, O(CO)R', OR', SR', COOR', R_{ph} , CR' = CR'- R_{ph} and CR_2' - CR_2' - R_{ph} (wherein R_{ph} represents an unsubstituted or substituted phenyl group with the phenyl substituents being chosen from any of the non-phenyl substituents defined for R^{17} - R^{20} and wherein R' is H or a lower alkyl group).

- 5. The compound of claim 1, structure A-E, wherein, Z = S, Y = N, R' = H, $R^1 = H$, $R^2 = CH_3$ and R^3 R^{14} are H.
- 6. The compound of claim 1, structure A-E, wherein, Z = S, Y = O, R' = H, $R^2 = CH_3$ and R^3 R^{14} are H.
- 7. The compound of claim 1, structure A-E, wherein Z = S, Y = N, R' = H, R^{1-} $^{4} = H$, $R^{5} = I$, and R^{6} R^{14} are H.
- 8. The compound of claim 1, structure A-E, wherein Z = S, Y = N, R' = H, R^{1-} $^4 = H$, $R^5 = I$, $R^8 = OH$ and $R^6 R^7$ and $R^9 R^{14}$ are H.
- 9. The compound of claim 1, structure A-E, wherein, Z = S, Y = N, R' = H, $R^1 = H$, $R^2 = CH_2-CH_2-CH_2-F$ and R^3-R^{14} are H.
- 10. The compound of claim 1, structure A-E, wherein, Z = S, Y = O, R' = H, $R^2 = CH_2$ -CH₂-F and R^3 R^{14} are H.
- 11. The compound of claim 1, structure A-E, wherein Z=S, Y=N, R'=H, R¹⁻⁷ = H, R⁸ \neq O-CH₂-CH₂-F and R⁹- R¹⁴ are H.
- 12. The compound of claim 1, structure A-E, wherein Z=S, Y=N, R'=H, $R^1 = CH_3$, $R^{2-7} = H$, $R^8 = O-CH_2-CH_2-F$ and R^9-R^{14} are H.





- The compound of claim 1, structure F-J, wherein, Z = S, Y = O, R' = H, 14. $R^2 = CH_3$ and R^3 - R^{14} are H.
- The compound of claim 1, structure F-J, wherein Z = S, Y = N, R' = H, R^{1-} 15. 4 = H, R 5 = I, and R 6 - R 14 are H.
- The compound of claim 1/2, structure F-J, wherein Z = S, Y = N, R' = H, R^{1-1} 16. 4 = H, R 5 = I, R 8 = OH and R 6 - R 7 /and R 9 - R 14 are H.
- The compound of claim/1, structure F-J, wherein, Z = S, Y = N, R' = H, $R^1 = H$, 17. $R^2 = CH_2-CH_2-CH_2-F$ and R^3-R^{14} are H.
- 18. The compound of claim 1, structure F-J, wherein, Z = S, Y = O, R' = H, $R^2 = CH_2-CH_2-F$ and $R^2 + R^{14}$ are H.
- The compound of claim 1, structure F-J, wherein Z = S, Y = N, R' = H, R^{1-} 19. 7 = H, R^{8} = O-CH₂-CH₂-F and R^{9} - R^{14} are H.
- The compound of claim 1, structure F-J, wherein Z = S, Y = N, R' = H, 20. $R^1 = CH_3$, $R^{2-7} = H$, $R^8 = O-CH_2-CH_2-F$ and R^9-R^{14} are H.
- The compound of claim 3, wherein at least one of the substituents R³ -R¹⁴ is 21. selected from the group consisting of CN, OCH3, OH and NH2.
- 22. The compound of claim 1, wherein the amyloid binding compound is selected from the group consisting of structure B, structure C and structure D; wherein $R^1 = H$, $R^2 = CH_3$ and R^8 is selected from the group consisting of CN, CH₃, OH, OCH₃ and NH₂.
- The compound of claim 22, wherein R^3 R^7 and R^9 R^{14} are H. 23.

- 24. The compound of claim 1, wherein the compound binds to A β with a dissociation constant (K_D) between 0.0001 and 10.0 μ M when measured by binding to synthetic A β peptide or Alzheimer's Disease brain tissue.
- 25. The compound of claim 3, wherein the compound binds to A β with a dissociation constant (K_D) between 0.0001 and 10.0 μ M when measured by binding to synthetic A β peptide or Alzheimer's Disease brain tissue.
- 26. A method for synthesizing a compound of claim 1 having at least one of the substituents R¹-R¹⁴ selected from the group consisting of ¹³¹I, ¹²⁵I, ¹²³I, ¹⁶Br, ¹⁵Br, ¹³F, and ¹³F, comprising the step of labeling a compound of claim 1 wherein at least one of the substituents R¹-R¹⁴ is a tri-alkyl tin, by reaction of the compound with a ¹³¹I, ¹²⁵I, ¹²³I, ¹⁶Br, ¹⁵Br, ¹³F, or ¹³F containing substance.
- 27. A method for synthesizing a compound of claim 1 having at least one of the substituents R^3 R^{14} selected from the group consisting of 131 I, 125 I, 123 I, 76 Br, 75 Br, 18 F, and 19 F, comprising the step of labeling a compound of claim 1, structures A-E or F-J, wherein Z = S, Y = N, $R^1 = H$ and at least one of the substituents R^3 - R^{14} is a tri-alkyl tin, by reaction of the compound with a 131 I, 125 I, 123 I, 76 Br, 75 Br, 18 F, or 19 F containing substance.
- 28. A pharmaceutical composition for *in vivo* imaging of amyloid deposits, comprising (a) a compound of claim 1 and (b) a pharmaceutically acceptable carrier.
- 29. A pharmaceutical composition for *in vivo* imaging of amyloid deposits, comprising (a) a compound of claim 1, structures A-E or F-J, wherein Z = S, Y = N, $R^1 = H$, and (b) a pharmaceutically acceptable carrier.
- 30. An *in vivo* method for detecting amyloid deposits in a subject, comprising the steps of:
- (a) administering a detectable quantity of the pharmaceutical composition of claim 28, and

- (b) detecting the binding of the compound to amyloid deposit in the subject.
- 31. The method of claim 30, wherein the amyloid deposit is located in the brain of a subject.
- 32. The method of claim 30, wherein the subject is suspected of having a disease or syndrome selected from the group consisting of Alzheimer's Disease, familial Alzheimer's Disease, Down's Syndrome and homozygotes for the apolipoprotein E4 allele.
- 33. The method of claim 30, wherein the detecting is selected from the group consisting of gamma imaging, magnetic resonance imaging and magnetic resonance spectroscopy.
- 34. The method of claim 33, wherein the detecting is done by gamma imaging, and the gamma imaging is either PET or SPECT.
- 35. The method of claim 30, wherein the pharmaceutical composition is administered by intravenous injection.
- 36. The method of claim 30, wherein the ratio of (i) binding of the compound to a brain area other than the cerebellum to (ii) binding of the compound to the cerebellum, in the subject, is compared to the ratio in normal subjects.
- 37. A method of detecting amyloid deposits in biopsy or post-mortem human or animal tissue comprising the steps of:
- (a) incubating formalin-fixed or fresh-frozen tissue with a solution of a compound of claim 1 to form a labeled deposit and then,
 - (b) defecting the labeled deposits.
- 38. The method of claim 37 wherein the solution is composed of 25-100% ethanol, with the remainder of the solution being water, wherein the solution is saturated with the compound having one of structures A-E or F-J.

- 39. The method of claim 37 wherein the solution is composed of an aqueous buffer containing 0-50% ethanol, wherein the solution contains 0.0001 to 100 μ M of the compound having one of structures A-E or F-J.
- 40. The method of claim 37 wherein the detecting is effected by microscopic techniques selected from the group consisting of bright-field, fluorescence, laser-confocal, and cross-polarization microscopy.
- 41. A method of quantifying the amount of amyloid in biopsy or post-mortem tissue comprising the steps of:
- a) incubating a radiolabeled derivative of a compound of claim 1 with a homogenate of biopsy or post-mortem tissue, wherein at least one of the substituents R¹-R¹⁴ of the compound is labeled with a radiolabel selected from the group consisting of ¹²⁵I, ³H, and a carbon-containing substituent as specified in claim 1, wherein at least one garbon is ¹⁴C,
- b) separating the tissue-bound from the tissue-unbound radiolabeled derivative of a compound of claim 1,
- c) quantifying the tissue-bound radiolabeled derivative of a compound of claim 1, and
- d) converting the units of tissue-bound radiolabeled derivative of a compound of claim 1 to units of micrograms of amyloid per 100 mg of tissue by comparison with a standard.
- 42. The method of claim 41, wherein the radiolabeled derivative is an amyloid binding compound having one of structures A-E or a water soluble, non-toxic salt thereof:

alp)

wherein Z is S, NR', O or CR' in which case the correct tautomeric form of the heterocyclic ring becomes an indole in which R' is H or a lower alkyl group:
-85-

is not a quaternary

wherein Y is NR^1R^2 , OR^2 , or SR^2 ;

wherein the nitrogen of

amine;

or an amyloid binding compound having one of structures F-J or a water soluble, non-toxic salt thereof:

Structure G

$$R_8$$
 R_9
 R_{10}
 R_{10}
 R_7

Structure H

Structure I

$$R_8$$
 Z
 Q
 R_{10}

or

Structure J

wherein each $\ensuremath{\Omega}$ is independently selected from one of the following structures:

Harman Street Comments of the Arman Street Comments of the Street Co



$$R_6$$
 R_5 $(CH_2)_n$ Wherein $n = 0, 1, 2, 3 \text{ or } 4,$ R_4 R_3

$$R_6$$
 R_5 Z R_4 U R_3

$$R_6$$
 R_5
 Z
 R_3
 Z
 R_3

$$R_6$$
 R_5
 R_3
 R_4
 R_3

$$\begin{array}{c|c}
R_6 & R_5 \\
\hline
Z & R_3 \\
\hline
R_4 & Z & R_3 \\
\hline
R_3 & R_3
\end{array}$$

wherein Z is S, NR', O, or C(R')2 in which R' is H or a lower alkyl group; wherein U is CR' (in which R' is H or a lower alkyl group) or N (except when U

$$R_6$$
 R_5 R_4 R_3);

= N, then Q is not

wherein Y is NR^1R^2 , OR^2 , or SR^2 ;

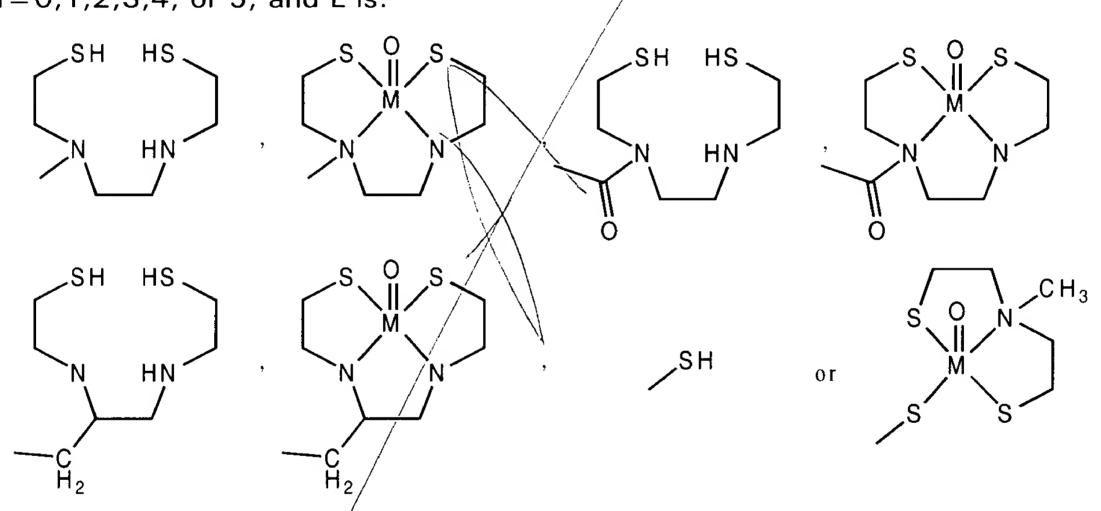
$$Z$$
or
 R_1

wherein the nitrogen of is not a quaternary or amine;

wherein each R¹ and R² independently is selected from the group consisting of H, a lower alkyl group, $(CH_2)_nOR'$ (wherein n = 1, 2, or 3), CF_3 , CH_2 - CH_2X , CH_2 -CH₂-CH₂X (wherein /X = F, CI, Br or I), (C = O)-R', R_{ph}, and (CH₂)_nR_{ph} (wherein n = 1, 2, 3, or 4 and Rph represents an unsubstituted or substituted phenyl group with the phenyl substituents being chosen from any of the non-phenyl substituents defined below for R3-R14 and R' is H or a lower alkyl group);



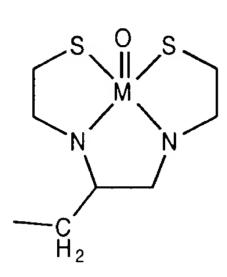
and wherein each R^3 - R^{14} independently is selected from the group consisting of H, F, CI, Br, I, a lower alkyl group, $(CH_2)_nOR'$ (wherein n=1, 2, or 3), CF_3 , CH_2 - CH_2X , O- CH_2 - CH_2X , CH_2 - CH_2 -C



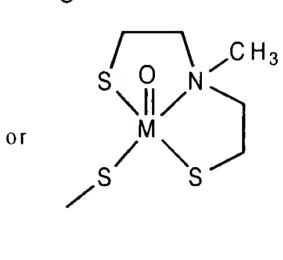
wherein M is selected from the group consisting of Tc and Re; or wherein each R^1 and R^2 is a chelating group (with or without a chelated metal group) of the form W-L, wherein W is $-(CH_2)_n$ where n=2,3,4, or 5; and L is:

HN

Atty. Dkt. No.: 076333-0281



, SH



wherein M is selected from the group consisting of Tc and Re;

or wherein each R^1-R^{14} independently is selected from the group consisting of a chelating group (with or without a chelated metal ion) of the form W-L and V-W-L, wherein V is selected from the group consisting of -COO-, and -CO-; W is - $(CH_2)_n$ where n=0,1,2,3,4, or 5; L is:

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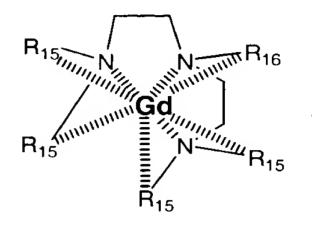
and wherein R¹⁵ independently is selected from the following:

H,
$$-COOH$$
, $-CONHCH_3$. $-CONHCH_3$. $-COOH$, $-CONHCH_3$. $-COOH$, $-COO$

or an amyloid binding, chelating compound (with or without a chelated metal group) or a water soluble, non-toxic salt thereof of the form:



$$R_{15}$$
 N N R_{16} R_{15}



wherein R¹⁵ independently is selected from the following:





and
$$R^{16}$$
 is H^{0} is H^{0} H

independently selected from one of the following structures:

R₁₇ R₁₈ wherein
$$n = 0, 1, 2, 3 \text{ or } 4,$$

R₁₇ R₁₈

R₁₇ R₁₈

R₁₇ R₁₈

R₁₈ R₁₉

R₁₇ R₁₈

R₁₇ R₁₈

R₁₇ R₁₈

R₁₈ R₁₉

R₁₉ R₁₉

R₁₉ R₁₉

wherein Z is S, NR', O, or C(R')2 in which R' is H or a lower alkyl group; wherein U is N or CR';

wherein Y is NR¹R², OR², or SR²;/

wherein each R^{17} - R^{24} independently is selected from the group consisting of H, F, CI, Br, I, a lower alkyl group, $(CH_2)_nOR'$ (wherein n=1, 2, or 3), CF_3 , CH_2 - CH_2X , O- CH_2 - CH_2X , CH_2 - CH_2



- 43. A method of distinguishing an Alzheimer's disease brain from a normal brain comprising the steps of:
- a) obtaining tissue from (i) the cerebellum and (ii) another area of the same brain other than the cerebellum, from normal subjects and from subjects suspected of having Alzheimer's disease;
- b) incubating the tissues with a radiolabeled derivative of a compound of claim 1 derivative so that amyloid in the tissue binds with the radiolabeled derivative of a compound of claim 1;
- c) quantifying the amount of amyloid bound to the radiolabeled derivative of a compound of claim 1, by administering a detectable quantity of the pharmaceutical composition comprising a compound of claim 1 with a pharmaceutically acceptable carrier, and detecting the binding of the compound to amyloid deposit in the subject;
- d) calculating the ratio of the amount of amyloid in the area of the brain other than the cerebellum to the amount of amyloid in the cerebellum;
- e) comparing the ratio for amount of amyloid in the tissue from normal subjects with ratio for amount of amyloid in tissue from subjects suspected of having Alzheimer's disease; and
- f) determining the presence of Alzheimer's disease if the ratio from the brain of a subject suspected of having Alzheimer's disease is above 90% of the ratios obtained from the brains of normal subjects.